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Article

Comparative Evaluations of Multiple Treatment Modalities for Meibomian Gland Dysfunction: A Prospective Clinical Study

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Abstract: Purpose: The aim of this study was to evaluate the different approaches used in the treatment of Meibomian gland dysfunction (MGD). Materials and Methods: The study included 92 cases with MGD who were assigned to treatment groups of conservative management alone or together with Intense Pulsed Light (IPL) therapy, oral azithromycin, oral doxycycline, topical cyclosporine and preservative-free artificial tears containing polyvinyl alcohol and povidone. Ocular surface parameters, including the Ocular Surface Disease Index (OSDI), tear break-up time (TBUT), SPEED score, and Meibomian gland parameters were evaluated. Results: A statistically significant increase was determined in Meibum grade plugging and Marx line scores with treatment of IPL therapy (p < 0.05). In the group receiving doxycycline treatment; there was a significant improvement in OSDI and SPEED scores, plugging, TBUT, and meibum grades. In the group receiving cyclosporine treatment, there was a significant improvement in OSDI and SPEED scores, plugging, TBUT, and meibum grades. There was a statistically significant increase in the conservative treatment group in respect of lid margin irregularity, Marx line score, OSDI and SPEED scores. In the group receiving azithromycin treatment, plugging, lid margin irregularity, TBUT, OSDI and SPEED scores increased significantly. Conclusions: Various treatment methods can be used for MGD. All the conservative treatment methods used in this study were seen to be successful.

Keywords: antibiotics; artificial tears; intense pulsed light therapy; meibomian gland dysfunction

1. Introduction

Meibomian gland dysfunction (MGD) constitutes the primary etiology of dry eye disease in clinical practice [1]. This condition, characterized as an evaporative form of dry eye, results from terminal duct obstruction of the Meibomian glands, leading to alterations in the lipid layer and lid margin integrity. Patients with MGD experience diminished meibum secretion with increased viscosity, and lid margin keratinization. The consequent tear film instability compromises ocular surface wetting, precipitating evaporative dry eye syndrome and triggering inflammatory processes that further exacerbate patient discomfort and visual disturbance [2,3]. Understanding these pathophysiological mechanisms is essential for the development of targeted therapeutic interventions which will address both the mechanical obstruction and inflammatory components of this prevalent condition.

Changes in meibum secretion can lead to increased colonization of micro-organisms along the lid margin, which may in turn contribute to infections [4]. The initial approach to treatment usually involves conservative measures such as applying warm compresses, performing eyelid massage, and maintaining eyelid hygiene with cleansers that include tea extract combined with the use of artificial tears [5,6]. However, it remains a subject of ongoing discussion whether the underlying cause of MGD is an infection at the lid margin or if the infection develops as a consequence of MGD [7].

Antibiotics play an important role in the treatment of both blepharitis and MGD because bacteria produce pro-inflammatory substances that can worsen the condition [8]. Tetracycline antibiotics (such as minocycline and doxycycline) are frequently used, as they not only inhibit bacterial growth but also reduce inflammation by suppressing bacterial lipase activity and controlling pro-inflammatory mediator production [9,10]. Azithromycin, a macrolide antibiotic, works in a similar way by limiting bacterial growth, reducing the release of lipase, and modulating pro-inflammatory molecules. It also helps to regulate the function and secretion of the Meibomian glands. Azithromycin can be administered both topically and orally [11,12].

Intense Pulsed Light (IPL) therapy offers another treatment modality. It utilizes a high-intensity, non-coherent light source (with wavelengths ranging from 500 to 1200 nm) that stimulates collagen production, targets and destroys abnormal blood vessels, and helps reduce the viscosity of meibum through heat. Furthermore, IPL has been shown to be effective against Demodex mites [13]. A further treatment option is Cyclosporine A, available as a 0.05% topical emulsion, which is a calcineurin inhibitor that was the first drug approved by the FDA for treating dry eye disease [6].

The aim of this study was to evaluate the effectiveness of various treatment approaches for MGD by assessing clinical parameters, to determine which treatment method is most effective and best enhances patient comfort.

2. Materials and Methods

The study included 92 cases diagnosed with MGD who presented to Kahramanmaraş Necip Fazıl City Hospital in February 2025. All the study procedures complied with the principles of the Helsinki Declaration. Approval for the study was granted by the Ethics Committee of Harran University (decision no: HRÜ/25.03.33). Informed consent was obtained from all the patients.

All the patients underwent a comprehensive ophthalmological assessment, comprising slit-lamp biomicroscopic examination, detailed evaluation of anterior segment structures and fundoscopic examination, determination of best-corrected visual acuity, and measurement of intraocular pressure using applanation tonometry.

Obstructive MGD was diagnosed based on the Japanese MGD diagnostic criteria [14]. The diagnosis was confirmed by the presence of ocular symptoms together with abnormal findings in the Meibomian gland orifices such as increased vascularity or the detection of meibomian gland orifice obstruction, as evidenced by plugging and reduced meibum secretion with the application of moderate digital pressure.

Exclusion criteria for the study were defined as signs of allergic or infectious conjunctivitis, significant refractive errors, ectatic corneal conditions, the presence of systemic diseases such as diabetes mellitus and hypertension, systemic rheumatological diseases, a history of prior ophthalmic surgery, or current contact lens usage. Care was taken to ensure that none of the patients had received any eye treatment in the last three months.

Following a comprehensive ophthalmologic evaluation, standardized assessments were made including the Ocular Surface Disease Index (OSDI) and the Standard Patient Evaluation for Eye Dryness (SPEED) scores. The Tear Break-Up Time (TBUT) test was applied and detailed lid margin evaluations were performed to assess ocular surface integrity. All patients were subsequently recalled for a one-month follow-up examination, during which the initial assessment protocols were repeated to monitor clinical response and treatment efficacy.

OSDI and **SPEED**

To quantitatively assess the impact of dry eye on quality of life, the 12-item Ocular Surface Disease Index (OSDI), and the 4-item Standard Patient Evaluation for Eye Dryness (SPEED) questionnaire, were administered to all the subjects (see Supplementary Table S1 and Figure 1).

TBUT

Tear film stability was evaluated using the standardized TBUT assessment. Following instillation of 1% fluorescein dye into the conjunctival sac, the time interval between the last complete blink and the first appearance of a corneal black spot in the stained tear film was measured under cobalt blue illumination. To ensure reliability, three consecutive measurements were obtained for each subject, and the mean value was calculated and recorded as the definitive TBUT

Grading of Lid Margin Findings and Meibum Expressibility

Lid margin findings were assessed with slit-lamp examination using standardized grading scales. Telangiectasia was evaluated on a scale of 0–3, where 0 indicates no findings, 1 represents mild telangiectasia, 2 corresponds to moderate telangiectasia, and 3 denotes severe telangiectasia. The mucocutaneous junction was similarly graded on a scale of 0–3.

For the Marx line (ML) score, 1% fluorescein drops were instilled on the lower eyelid, and the patient was instructed to blink several times. The lower eyelid was then partitioned into three regions — outer, middle, and inner — and points were assigned based on the extent of fluorescein contact with the Meibomian orifices (MOs). Specifically, an ML score of 0 was assigned when the Marx line courses on the skin side of the MO line without contacting the orifices, 1 when portions of the Marx line touch the MOs, 2 when the Marx line passes directly through the MOs, and 3 when the Marx line courses along the eyelid margin side of the MOs. Lid irregularity, plugging, and foaming were each graded on a scale of 0–2, where 0 signifies no findings, 1 reflects mild findings, and 2 indicates severe findings [14,15].

The expressibility of meibum from the central area of both the upper and lower eyelids was evaluated semi-quantitatively using a scale of 0–3: 0 indicated that clear meibum was readily expressed; 1 that cloudy meibum was expressed with mild pressure; 2 that cloudy meibum required more than moderate pressure for expression; and 3 that meibum could not be expressed even under strong pressure [16,17].

IPL

For the IPL procedure, the Eye-light® device (Topcon) was utilized. Initially, 0.5% proparacaine anesthetic eye drops were instilled in both eyes. The treatment involved administering a series of 20 overlapping light pulses to the targeted periorbital skin, extending from the right temple across the lower eyelid, including the nasal bridge, and continuing to the left temple along the lower eyelid. A 590 nm filter in combination with a 6 mm cylindrical light guide was employed on the handpiece.

Treatment Groups for MGD Patients

The patients diagnosed with MGD were stratified into five distinct groups based on the treatment modalities, all of which were applied for one month.

Group 1 (Only Conservative Treatment)

In Group 1, patients received conservative management only. All the subjects were prescribed preservative-free artificial tears (4 times daily) containing polyvinyl alcohol and povidone (Novaqua, DEVA Holding Inc., Turkey). A comprehensive conservative regimen was recommended, which included the application of warm compresses, performing eyelid massage prior to bedtime, and cleansing the eyelashes with an eyelash shampoo.

Group 2 (Conservative Management with Adjunctive Oral Doxycycline Therapy)

In Group 2, patients were administered oral doxycycline 100 mg (Monodox, DEVA Holding Inc., Turkey) as an adjunct to the conservative management regimen. The dosage was structured with twice-daily administration during the initial week, then once daily for the subsequent three weeks, providing a treatment duration of one month.

Group 3 (Conservative Management with Adjunctive Oral Azithromycin Therapy)

In Group 3, oral azithromycin 500 mg (Azitro, DEVA Holding Inc., Turkey, 500 mg) was administered as a single daily dose for three consecutive days in conjunction with the conservative treatment regimen.

Group 4 (Conservative Management with Adjunctive Topical Cyclosporine Treatment)

In Group 4, topical cyclosporine 0.05% ophthalmic emulsion (Ocurin, Bilim Pharmaceuticals, Turkey) was prescribed as an adjunctive therapy to the conservative treatment protocol. The cyclosporine drops were administered twice daily.

Group 5 (Conservative Management with Adjunctive IPL Therapy)

Together with the conservative management, the IPL protocol was implemented on days 0, 15, and 30.

Statistical Analyses

Data obtained in the study were analyzed statistically using SPSS version 27 software. The suitability of the variables to normal distribution was examined with Kolmogorov-Smirnov/Shapiro Wilk tests. Descriptive statistics were expressed as mean±standard deviation (SD) or median values for continuous data and as number (n) and percentage (%) for categorical data. For values not showing normal distribution, the Wilcoxon test was applied. A value of p<0.05 was considered statistically significant.

3. Results

The 92 subjects enrolled in this study, comprised 50 (64.2%) females and 40 (35.8%) males with a mean age of 45.16 ± 7.28 years.

Group 1 (conservative treatment only) included 18 subjects (6 males, 12 females) with a mean age of 46.67 years. Group 2 (doxycycline) included 22 subjects (8 males, 14 females) with a mean age of 45.38 years. Group 3 (azithromycin) included 18 subjects (6 males, 12 females) with a mean age of 46.44 years. Group 4 (cyclosporine) included 18 subjects (5 males, 13 females) with a mean age of 47.11 years. Group 5 (IPL) included 16 subjects (5 males, 11 females) with a mean age of 43.88 years. No statistically significant differences were observed in age and gender distributions among the treatment groups (p>0.05).

Group 1: The TBUT values increased slightly from 9.00 ± 2.67 seconds to 9.94 ± 2.41 seconds. The lid vascularity scores decreased significantly from 1.67 ± 0.84 to 1.06 ± 0.80 (p=0.014). Meibomian gland plugging was determined to have improved slightly from 1.11 ± 0.58 to 0.94 ± 0.41 . Lid margin irregularity demonstrated a significant reduction from 0.67 ± 0.48 to 0.33 ± 0.48 (p=0.014). The foaming scores decreased from 0.89 ± 1.56 to 0.44 ± 0.51 . Marx line scores showed a significant improvement from 4.28 ± 0.89 to 3.33 ± 1.18 (p=0.003). The meibum grade improved slightly from 1.67 ± 0.48 to 1.50 ± 0.51 . Patient-reported outcomes were seen to be significantly improved, with OSDI scores decreasing from 51.67 ± 45.50 to 38.89 ± 8.83 (p<0.01), and SPEED scores reducing from 0.46 ± 0.15 to 0.29 ± 0.12 (p<0.01).

Group 2: The TBUT values increased significantly from 8.15 ± 3.23 to 10.58 ± 3.11 seconds (p<0.01). a statistically significant decrease was determined in the lid vascularity values from 1.46 ± 0.70 to 2.08 ± 0.93 (p<0.01), and in the Meibomian gland plugging values from 1.54 ± 0.76 to 1.12 ± 0.43 (p<0.01). Lid margin irregularity improved from 0.85 ± 0.50 to 0.58 ± 0.50 , foaming decreased from 0.85 ± 0.96 to 0.58 ± 0.75 , and the Marx line scores improved from 5.46 ± 1.74 to 4.62 ± 1.09 . Statistically significant improvements were seen in Meibum grade from 2.00 ± 0.66 to 1.54 ± 0.58 (p<0.01), in OSDI from 46.77 ± 16.22 to 33.77 ± 8.37 (p<0.01) and in SPEED from 0.55 ± 0.14 to 0.34 ± 0.18 (p<0.01).

Group 3: A statistically significant increase was determined in TBUT from 6.83 ± 1.24 to 9.00 ± 1.28 seconds (p<0.01). Lid vascularity decreased from 2.00 ± 0.97 to 1.33 ± 0.68 (p=0.007), and Meibomian gland plugging from 1.44 ± 0.70 to 1.00 (p=0.023). Lid margin irregularity improved from 1.11 ± 0.32 to

 0.89 ± 0.58 (p=0.046), foaming decreased from 0.89 ± 0.58 to 0.56 ± 0.51 (p=0.00), and the Marx line score showed a slight improvement from 5.89 ± 1.45 to 5.50 ± 1.65 . The Meibum grade improved from 1.89 ± 0.58 to 1.44 ± 0.51 . statistically significant improvements were seen in the patient-reported outcomes, with a decrease in OSDI from 41.11 ± 18.69 to 27.33 ± 7.91 (p=0.003) and in SPEED from 0.48 ± 0.21 to 0.32 ± 0.17 (p=0.002).

Group 4: The decrease in TBUT from 11 ± 1.32 to 9.17 ± 0.98 seconds was statistically significant (p<0.01). Lid vascularity showed significant improvement from 1.78 ± 0.94 to 1.28 ± 0.75 (p=0.014). Meibomian gland plugging significantly reduced from 1.11 ± 0.32 to 0.89 ± 0.58 (p=0.046), lid margin irregularity slightly increased from 0.33 ± 0.33 to 0.44 ± 0.51 , and foaming remained stable at 0.33 ± 0.48 . The Marx line score showed a minimal improvement from 4.00 ± 0.84 to 3.89 ± 1.27 . A significant improvement was observed in the Meibum grade from 1.78 ± 0.64 to 1.44 ± 0.70 (p=0.014). Patient-reported outcomes showed significant improvements, with a decrease in OSDI from 52.33 ± 14.66 to 40.33 ± 9.44 (p<0.01) and in SPEED from 0.46 ± 0.15 to 0.29 ± 0.12 (p<0.01).

Group 5: A statistically significant increease was determined in TBUT from 7.20 \pm 1.13 seconds to 10.40 \pm 0.51 seconds (p=0.004). Lid vascularity scores decreased from 2.00 to 1.33 \pm 0.97, and Meibomian gland plugging reduced from 1.40 \pm 0.51 to 1.00 \pm 0.94 (p=0.046). Pre and post-treatment lid margin irregularity remained stable at 0.80 \pm 0.78, and foaming scores were maintained at 0.20 \pm 0.42. A statistically significant improvement was observed in the Marx line scores declining from 4.80 \pm 1.22 to 3.60 \pm 0.51 (p=0.014), and in Meibum grade decreasing from 2.00 \pm 0.66 to 1.60 \pm 0.84 (p=0.046). Patient-reported outcomes also improved substantially, with a decrease in OSDI scores from 47.20 \pm 20.06 to 19.60 \pm 7.00 and in SPEED from 0.62 \pm 0.13 to 0.34 \pm 0.07.

3.1. Figures, Tables and Schemes

S	PEED"	M QUE	STION	NAIRE		
Name:	Date	e://	Sex: N	/ F (Circle)	DOB:/_	_/
For the Standardized Patient Evaluatic checking the box that best represents					the following qu	iestions by
1. Report the type of <u>SYMPTOMS</u> yo	u experience a	and when the	ey occur:			
	At this	visit	Within past	72 hours	Within past 3	months
Symptoms	Yes	No	Yes	No	Yes	No
Dryness, Grittiness or Scratchiness						
Soreness or Irritation						
Burning or Watering						
Eye Fatigue						
2. Report the <u>FREQUENCY</u> of your sy Symptoms	mptoms using	the rating li	st below:	3		
Dryness, Grittiness or Scratchiness					1	
Soreness or Irritation					1	
					-	
Burning or Watering Eye Fatigue					-	
3. Report the <u>SEVERITY</u> of your symp	toms using th	e rating list b	elow:			
Symptoms	0	1	2	3	4	
Dryness, Grittiness or Scratchiness						
Soreness or Irritation						
Burning or Watering						
Eye Fatigue						
0 = No Problems 1 = Tolerable - not perfect, but not uncom 2 = Uncomfortable - irritating, but does n 3 = Bothersome - irritating and interferes 4 = Intolerable - unable to perform my da 4. Do you use eye drops for lubrication	ot interfere with with my day ily tasks	my day	IO If yes, ho	ow often?		
Cornea. 2013 Sep;32(9):1204-10 © 2011 TearScience, Inc. All rights reserved. 13-ADV-123 A				office use only	requency + Seve	
	M)P			

Figure 1. Speed Questionnaire.

Table 1. Comparisons of Pre-treatment and Post-treatment Clinical Parameters in Patients with Meibomian Gland Dysfunction Across Different Treatments.

	Group 1-		Group 2-		Group 3-		Group 4-		Group 5-	
	conservati	We	conservati	We.	conservati	ive	conservative		conservative	
	treatment		treatment		treatment		treatment		treatment + IPL	
	treatment	Offry	doxycyclii		azithromy				therapy	
		I	doxycyciii	1	uzitiioiity	- Ciri	cycloopor		uncrup)	
Clinical	Pre-	Post-	Pre-	Post-	Pre-	Post-	Pre-	Post-	Pre-	Post-
Paramet	treatme	treatme	treatme	treatme	treatme	treatme	treatme	treatme	treatme	treatme
ers	nt	nt	nt	nt	nt	nt	nt	nt	nt	nt
	(mean±s	(mean±	(mean±s	(mean±	(mean±s	(mean±	(mean±s	(mean±	(mean±s	(mean±
	d)	sd)	d)	sd)	d)	sd)	d)	sd)	d)	sd)
Lid	1.67±0.8	1.06±0.8	1.46±0.7	2.08±	2.00±0.9	1.33±0.6	1.78±0.9	1.28±0.7	2.00±0.9	1.33±0.9
vascular	4	0	0	0.93	7	8	4	5	7	7
ity										
Pluggin	1.11±0.5	0.94±0.4	1.54±0.7	1.12±0.4	1.44±0.7	1.00±0.0	1.11±0.3	0.89±0.5	1.40±0.5	1.00±0.9
g	8	1	6	3	0	0	2	8	1	4
Lid	0.67±0.4	0.33±0.4	0.85±0.5	0.58±0.5	1.11±0.3	0.89±0.5	0.33±0.3	0.44±0.5	0.80±0.7	0.80±0.7
margin	8	8	0	0	2	8	3	1	8	8
irregula										
rity										
Foamin	0.89±1.5	0.44±0.5	0.85±0.9	0.58±0.7	0.89±0.5	0.56±0.5	0.33±0.4	0.33±0.4	0.20±0.4	0.20±0.4
g	6	1	6	5	8	1	8	8	2	2
Marx	4.28±0.8	3.33±1.1	5.46±1.7	4.62±1.0	5.89±1.4	5.50±1.6	4.00±0.8	3.89±1.2	4.80±1.2	3.60±0.5
line	9	8	4	9	5	5	4	7	2	1
score										
Meibum	1.67±0.4	1.50±0.5	2.38±0.4	1.54±0.5	1.89±0.5	1.44±0.5	1.78±0.6	1.44±0.7	2.00±0.6	1.60±0.8
grade	8	1	6	8	8	1	4	0	6	4
OSDI	51.67±45	38.89±8.	46.77±16	33.77±8.	41.11±18	27.33±7.	52.33±14	40.33±9.	47.20±20	19.60±7
	.50	83	.22	37	.69	91	.66	44	.06	

Speed	0.46±0.1	0.29±0.1	0.55±0.1	0.34±0.1	0.48±0.2	0.32±0.1	0.54±0.1	0.37±0.1	0.62±0.1	0.34±0.0
	5	2	4	8	1	7	5	1	3	7
TBUT	9.00±2.6	9.94±2.4	8.15±3.2	10.58±3.	6.83±1.2	9.00±1.2	7.11±1.3	9.17±0.9	7.20±1.1	10.40±0.
	7	1	3	11	4	8	2	8	3	51

Γable 2. Grading scales for the evalua	ation criteria of Meibomian gland dysfunction.
Lid vascularity	
0	None
0	
1	Redness of the palpebral conjunctiva with no vascularity
	around the gland orifices
2	Redness of the palpebral conjunctiva with vascularity
	around the gland orifices affecting<50% of the full length of
	the lid margin
3	Redness of the palpebral conjunctiva with vascularity
	around the gland orifices affecting≥50% of the full length of
	the lid margin
Lid margin irregularity	
0	None
1	Fewer than three lid margin irregularities with shallow
	notching
2	Three or more lid margin irregularities or deep notching
Foaming	
0	None
1	Mild findings
2	Severe findings
Lid plugging	
0	None
1	Plugging of <3 gland orifices
2	Plugging of ≥3 gland orifices affecting<50% of the full
	length of the lid margin
3	Plugging of ≥3 gland orifices affecting≥50% of the full
	length of the lid margin

Marx line score	
0	Marx line running entirely along the conjunctival side of
	the gland orifices
1	Part of the Marx line in contact with the gland orifices
2	Marx line running through the gland orifices
3	Marx line running along the eyelid margin on the side of
	the gland orifices
Meibum grade	
0	Clear meibum is easily expressed
1	Cloudy meibum is expressed with mild pressure
2	Cloudy meibum is expressed with more than moderate
	pressure
3	Meibum cannot be expressed even with the hard pressure

4. Discussion

MGD is an increasingly prevalent ocular surface disorder, characterized by terminal duct obstruction and/or qualitative/quantitative changes in glandular secretion. This results in altered tear film lipid layers, evaporative dry eye, and chronic ocular surface inflammation. The multifactorial nature of MGD calls for equally diverse treatment approaches [20,21]. It was observed in the literature that while several studies have compared oral treatment modalities for MGD, no comparative studies have specifically evaluated the effectiveness of cyclosporine versus IPL therapy.

Conservative management (preservative-free artificial tears, warm compresses, eyelid massage, and eyelash cleansing) forms the foundation of MGD treatment, providing relief for mild cases by improving meibum properties and tear film stability. However, this approach alone often fails to address underlying glandular abnormalities in moderate-to-severe cases or those with persistent inflammation, necessitating adjunctive therapies to target the complex pathophysiology of MGD.

In Groups 2 and 3, adjunctive oral antibiotic therapy was evaluated. Doxycycline, a tetracycline antibiotic, exerts anti-inflammatory effects by inhibiting matrix metalloproteinases and modifying the lipid composition of meibum, which together reduce inflammation and enhance glandular function. In contrast, oral azithromycin provides the advantage of a shorter treatment course, improved patient compliance, and a lower incidence of adverse effects. The clinical distinction between these antibiotics is significant, as effective adjunctive antibiotic therapy hinges on both antimicrobial activity and the modulation of inflammatory pathways in MGD. The current study findings support previous studies that have reported that both treatments lead to improved TBUT, reduced meibomian gland plugging, and better subjective patient outcomes. There was also observed to be a greater improvement in eyelid symptoms with azithromycin than with doxycycline. Kashkouli et al. and Bukhari et al. compared oral azithromycin and doxycycline treatments, and reported that azithromycin was more effective on conjunctival redness and corneal staining [22,23]. In another study comparing topical azithromycin and oral doxycycline treatment, oral doxycycline treatment caused a greater prolongation of tear break-up time, whereas topical azithromycin treatment had a greater effect on symptoms [24].

Topical cyclosporine was incorporated in the treatment of Group 4 in this study, reflecting an alternative therapeutic approach that leverages immunomodulation. Topical cyclosporine is known to suppress T-cell activity and reduce ocular surface inflammation [6,25]. Although primarily used in the management of chronic dry eye syndrome, its role in MGD is gaining recognition, particularly in cases where inflammation is a dominant feature. The statistically significant improvement in lid

vascularity and meibum quality observed in this group suggests that localized immunomodulatory therapy can be effectively integrated into a comprehensive treatment regimen for MGD. These results are encouraging, especially for patients who might be contraindicated for systemic therapy or who experience adverse effects related to systemic antibiotics. Jeon et al. found that patients treated with a combination of cyclosporine and IPL showed reduced inflammatory markers and improved eyelid condition. These results hint at a possible synergistic effect between the two treatments, tackling inflammation and enhancing clinical signs in a complementary manner [26]. Iaccheri et al. applied only artificial tear drop treatment to one group of patients with MGD-related dry eye and artificial tear drop treatment together with cyclosporine drops to the other group, and observed a significant improvement in meibum expression and quality from the first months in the group in which cyclosporine treatment was added [27].

In Group 5 of the current study, which received adjunctive IPL therapy, significant clinical improvements were observed one month post-treatment. The thermal effects of IPL appear to liquefy meibum and reduce abnormal telangiectatic vessels, thereby modulating inflammatory mediators and enhancing the ocular surface microenvironment. In this study, IPL treatment was associated with decreased meibomian gland plugging, improved Marx line scores and meibum grade, and a longer TBUT. These results align with previous findings that support the efficacy of IPL in reducing MGD symptoms and improving both eyelid and gland function [21].

Based on the pre- and post-treatment score reductions, slight improvements were seen in Group 1, which served as the baseline. In the comparisons, greater reductions in parameters related to tear film stability and gland obstruction were observed in Groups 2 and 3, reflecting the benefits of systemic anti-inflammatory and lipid-modulating effects. Significant improvements in inflammatory indicators and lid vascularity were determined in Group 4, highlighting the advantage of localized immunomodulation, and inflammatory markers without systemic effects, making this treatment suitable for patients with contraindications to oral antibiotics. The most significant overall score reductions especially in meibomian gland plugging, Marx line scores, and TBUT were obtained in Group 5, indicating that IPL therapy was more effective on both the obstructive and inflammatory components of MGD. While conservative management provides the basis of MGD treatment, adjunctive therapies can significantly enhance outcomes. Systemic antibiotics and topical cyclosporine were seen to improve tear film stability and reduce inflammation, whereas IPL therapy emerged as the most effective non-invasive modality for comprehensive management due to its superior impact on both obstructive and inflammatory components.

Limitations

As topical azithromycin is not available in Türkiye, this treatment modality could not be included, so no comparison could be made with oral azithromycin. A further limitation could be said to be that the one-month follow-up period may not have been sufficient to fully capture the long-term benefits of treatments such as topical cyclosporine, which has been shown to be more effective with extended use. Variations in follow-up periods in the literature also pose challenges for direct comparison of the current study findings with those of previous studies.

5. Conclusions

The results of this study demonstrate that while conservative management provides a baseline therapeutic effect in MGD, adjunctive therapies significantly enhance clinical outcomes. Oral antibiotics (doxycycline and azithromycin) improved tear film stability and reduced glandular obstruction, while topical cyclosporine effectively targeted ocular surface inflammation. Notably, IPL therapy emerged as the most efficacious adjunctive modality, providing greater improvements in both objective clinical parameters and subjective patient-reported outcomes. These findings support a personalized, multimodal approach to MGD management, with treatment selection guided by individual patient characteristics and disease severity. Further research with extended follow-up periods is warranted to establish long-term efficacy profiles of these interventions

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Appendix A

Supplementary Table 1

Table S1. This is a table caption. Supplementary Table 1. OSDI.

Have you experienced any of the following during the last week:								
	All of the time	Most of the time	Half of the time	Some of the time	None of the time			
1. Eyes that are sensitive to light	4	3	2	1	0			
2. Eyes that feel gritty?	4	3	2	1	0			
3. Painful or sore eyes?	4	3	2	1	0			
4. Blurred vision?	4	3	2	1	0			
5. Poor vision?	4	3	2	1	0			
Have problems with your eyes limited you in performing any of the following during the last week:								
	All of the time	Most of the time	Half of the time	Some of the time	None of the time			
6. Reading?	4	3	2	1	0	N/A		

7. Driving at night?	4	3	2	1	0	N/A
8. Working with a computer or a bank machine(ATM)?	4	3	2	1	0	N/A
9. Watching TV?	4	3	2	1	0	N/A

Have your eyes felt uncomfortable in any of the following situations during the last week:

	All of the time	Most of the time	Half of the time	Some of the time	None of the time	
10. Windy conditions?	4	3	2	1	0	N/A
11. Places or areas with low humidity (very dry)?	4	3	2	1	0	N/A
12. Areas that are air conditioned?	4	3	2	1	0	N/A

Appendix B

Figure 1

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